

Antimicrobial Resistance Reporting: CP-CRE and *Candida auris*

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Antimicrobial Resistant Reportable Diseases

- Carbapenem-producing, carbapenem-resistant *Enterobacteriaceae* (CP-CRE)
- *Candida auris*
- Vancomycin Intermediate/Resistant *Staphylococcus aureus* (VISA/VRSA)
- Unusual occurrence, outbreak, or epidemic

2019 REPORTABLE DISEASES IN MICHIGAN – BY CONDITION

A Guide for Physicians, Health Care Providers and Laboratories

Report the following conditions to the Michigan Disease Surveillance System (MDSS) or local health department (see reverse) within 24 hours (unless otherwise noted) if the agent is identified by clinical or laboratory diagnosis.

Report the unusual occurrence, outbreak or epidemic of any disease or condition, including healthcare-associated infections.

Acute flaccid myelitis (1)	Listeriosis (<i>Listeria monocytogenes</i>) (5,6)
Anaplasmosis (<i>Anaplasma phagocytophilum</i>)	Lyme Disease (<i>Borrelia burgdorferi</i>)
Anthrax (<i>Bacillus anthracis</i> and <i>B. cereus</i> serovar anthracis) (4)	Malaria (<i>Plasmodium</i> species)
Arboviral encephalitides, neuro- and non-neuroinvasive: Chikungunya, Eastern Equine, Jamestown Canyon, La Crosse, Powassan, St. Louis, West Nile, Western Equine, Zika (6)	Measles (Measles/Rubeola virus)
Babesiosis (<i>Babesia microti</i>)	Melioidosis (<i>Burkholderia pseudomallei</i>) (4)
Blastomycosis (<i>Blastomyces dermatitidis</i>)	Meningitis: bacterial, viral, fungal, parasitic and amebic
Botulism (<i>Clostridium botulinum</i>) (4)	Meningococcal Disease (<i>Neisseria meningitidis</i> , sterile sites) (5)
Brucellosis (<i>Brucella</i> species) (4)	Middle East Respiratory Syndrome (MERS-CoV) (5)
Campylobacteriosis (<i>Campylobacter</i> species)	Mumps (Mumps virus)
Candidiasis (<i>Candida auris</i>) (4)	Orthopox viruses, including: Smallpox, Monkeypox (4)
Carbapenemase Producing – Carbapenem Resistant Enterobacteriaceae (CP-CRE): <i>Klebsiella</i> spp., <i>Enterobacter</i> spp., and <i>Escherichia coli</i> (5)	Pertussis (<i>Bordetella pertussis</i>)
Chancroid (<i>Haemophilus ducreyi</i>)	Plague (<i>Yersinia pestis</i>) (4)
Chickenpox / Varicella (<i>Varicella-zoster</i> virus) (6)	Polio (Poliovirus)
Chlamydial infections (including trachoma, genital infections, LGV) (<i>Chlamydia trachomatis</i>) (3, 6)	Prion disease, including CJD
Cholera (<i>Vibrio cholera</i>) (4)	Psittacosis (<i>Chlamydia psittaci</i>)
Coccidioidomycosis (<i>Coccidioides immitis</i>)	Q Fever (<i>Coxiella burnetii</i>) (4)
Cryptosporidiosis (<i>Cryptosporidium</i> species)	Rabies (<i>Rabies virus</i>) (4)
Cyclosporiasis (<i>Cyclospora</i> species) (5)	Rabies: potential exposure and post exposure prophylaxis (PEP)
Dengue Fever (Dengue virus)	Rubella (<i>Rubella virus</i>) (6)
Diphtheria (<i>Corynebacterium diphtheriae</i>) (5)	Salmonellosis (<i>Salmonella</i> species) (5)
Ehrlichiosis (<i>Ehrlichia</i> species)	Severe Acute Respiratory Syndrome (SARS) (5)
Encephalitis, viral or unspecified	Shigellosis (<i>Shigella</i> species) (5)
<i>Escherichia coli</i> , O157:H7 and all other Shiga toxin positive serotypes (5)	Spotted Fever (<i>Rickettsia</i> species)
Giardiasis (<i>Giardia</i> species)	<i>Staphylococcus aureus</i> , vancomycin intermediate/ resistant (VISA) (5)/VRSA (4))
Glanders (<i>Burkholderia mallei</i>) (4)	<i>Streptococcus pneumoniae</i> , sterile sites
Gonorrhea (<i>Neisseria gonorrhoeae</i>) (3, 6)	<i>Streptococcus pyogenes</i> , group A, sterile sites, including <i>Streptococcal Toxic Shock Syndrome</i> (STSS)
Guillain-Barre Syndrome (1)	Syphilis (<i>Treponema pallidum</i>) (6)
<i>Haemophilus influenzae</i> , sterile sites only- submit isolates for serotyping for patients < 15 years of age (5)	Tetanus (<i>Clostridium tetani</i>)
Hantavirus	Toxic Shock Syndrome (non-streptococcal) (1)
Hemolytic Uremic Syndrome (HUS)	Trichinellosis (<i>Trichinella spiralis</i>)
Hemorrhagic Fever Viruses (4)	Tuberculosis (<i>Mycobacterium tuberculosis</i> complex); report preliminary and final rapid test and culture results (4)
Hepatitis A virus (Anti-HAV IgM, HAV genotype)	Tularemia (<i>Francisella tularensis</i>) (4)
Hepatitis B virus (HBsAg, HBeAg, anti-HBc IgM, HBV NAAT, HBV genotype; report all HBsAg and anti-HBs (positive, negative, indeterminate) for children ≤ 5 years of age) (6)	Typhoid Fever (<i>Salmonella typhi</i>) and Paratyphoid Fever (serotypes Paratyphi A, Paratyphi B (tartrate negative), and Paratyphi C) (5)
Hepatitis C virus (all HCV test results including positive and negative antibody, RNA, and genotype tests) (6)	Vibriosis (Non-cholera vibrio species) (5)
Histoplasmosis (<i>Histoplasma capsulatum</i>)	Yellow Fever (Yellow Fever virus)
HIV (tests including reactive immunoassays (e.g., Ab/Ag, TD1/TD2, WB, EIA, IA), detection tests (e.g., VL, NAAT, p24, genotypes), CD4 counts/percents, and all tests related to perinatal exposures) (2,6)	Yersiniosis (<i>Yersinia enterocolitica</i>)
Influenza virus (weekly aggregate counts) Pediatric influenza mortality, report individual cases (5) Novel influenza viruses, report individual cases (5,6)	
Kawasaki Disease (1)	
Legionellosis (<i>Legionella</i> species) (5)	
Leprosy or Hansen's Disease (<i>Mycobacterium leprae</i>)	
Leptospirosis (<i>Leptospira</i> species)	

LEGEND

- (1) Reporting within 3 days is required.
 - (2) Reporting within 7 days is required.
 - (3) Sexually transmitted infection for which expedited partner therapy is authorized. See www.michigan.gov/hivstd for details.
 - (4) A laboratory shall immediately submit suspect or confirmed isolates, subcultures, or specimens from the patient being tested to the MDHHS Lansing laboratory.
 - (5) Isolate requested. Enteric: If an isolate is not available from non-culture based testing, the positive broth and/or stool in transport medium must be submitted to the MDHHS Lansing laboratory. Respiratory: Submit specimens, if available.
 - (6) Report pregnancy status, if available.
- Blue Bold Text = Category A bioterrorism or select agent, notify the MDHHS Laboratory immediately: (517) 335-8063

**Carbapenemase-producing
Carbapenem-resistant
Enterobacteriaceae (CP-CRE)**



What are CP-CRE?

- ***Enterobacteriaceae*** – enteric bacteria, gram negative bacilli
- **Carbapenems** – class of broad-spectrum, β -lactam antibiotics
 - Used for the treatment of resistant infections
 - Only 4 carbapenems: doripenem, ertapenem, imipenem, and meropenem
- **Carbapenemases** – enzymes that break down carbapenem antibiotics
 - Often encoded on plasmids which can be shared between different bacteria

Risk Factors for CP-CRE



Older age



**Antibiotic
use**



**Multiple
healthcare
exposures**

Acute care
Long-term care
Outside US
Recent surgery



**Indwelling
devices**

Catheters
Ventilators
PEG tubes



**Comorbid
conditions**

Cardiovascular
disease
Diabetes
Chronic lung
disease
Renal disease

CP-CRE Reporting

- Reportable disease in Michigan starting January 2018
- Surveillance definition endorsed by CSTE/CDC, nationally notifiable
- CP-CRE cases are reported using the Michigan Disease Surveillance System (MDSS)
 - Web-based communicable disease reporting system for the state of Michigan
 - Cases can be reported by:
 - Electronic laboratory report (ELR)
 - Manual case entry

CP-CRE Reporting Requirements

- Laboratories, infection prevention and Local Health Departments are required to report all cases of **CP-CRE** according to the following criterion for *Klebsiella spp.*, *E. coli*, or *Enterobacter spp.*:
 - Healthcare record contains a diagnosis of Carbapenemase-producing Carbapenem-resistant Enterobacteriaceae (CP-CRE), KPC, NDM, OXA-48, IMP or VIM or other novel carbapenemase
 - Any isolate of *Klebsiella spp.*, *E. coli*, or *Enterobacter spp.* demonstrating carbapenemase production by a phenotypic test (e.g., Carba NP, CIM, mCIM)
 - Any isolate of *Klebsiella spp.*, *E. coli*, or *Enterobacter spp.* with a known carbapenemase resistance mechanism (e.g., KPC, NDM, OXA-48, IMP, VIM, or other carbapenemase gene) by a recognized molecular test (e.g., PCR, Expert Carba-R)

CP-CRE Reporting Requirements

- If laboratories are unable to detect CP-CRE, (*i.e.*, cannot test for carbapenemase production (phenotypic) or resistance mechanism (molecular test):
 - Report any isolate of *Klebsiella spp.*, *E. coli*, or *Enterobacter spp.* with a minimum inhibitory concentration (MIC) of any of the following:
 - ≥ 4 mcg/ml for Meropenem
 - ≥ 4 mcg/ml for Imipenem
 - ≥ 4 mcg/ml for Doripenem
 - ≥ 2 mcg/ml for Ertapenem

Carbapenemase and Resistance Mechanism Testing

- Laboratories are *strongly encouraged to submit CRE isolates* to the MDHHS Bureau of Laboratories
 - Confirm organism identification
 - Perform modified carbapenem inactivation method (mCIM) testing
 - Perform PCR testing for KPC, NDM, OXA-48 like, IMP, VIM
 - If mCIM or PCR are positive, antimicrobial susceptibility testing (AST) will be performed

MDHHS Bureau of Laboratories Report

Antimicrobial Resistance Confirmation (ARC)

Gram Stain

Gram negative bacilli

Culture Results

Confirmed as *Klebsiella pneumoniae*

Identification Performed by MALDI-TOF.

Antimicrobial Susceptibility Results

	Klebsiella pneumoniae	
	MIC - Interpretation	
Amikacin	<=4	S
Aztreonam	>16	R
Cefepime	4	SDD
Cefotaxime	32	R
Ceftazidime	>16	R

Modified Carbapenem Inactivation Method
Positive

Phenotypic test

Modified Carbapenem Inactivation Method (mCIM) screen positive - this isolate demonstrates carbapenemase production. The clinical efficacy of the carbapenems has not been established for treating infections caused by Enterobacteriaceae and Pseudomonas aeruginosa that test carbapenem susceptible but demonstrate carbapenemase production in vitro. ISOLATES THAT ARE mCIM POSITIVE SHOULD BE CONSIDERED RESISTANT TO ALL CARBAPENEMS REGARDLESS OF MIC. MIC REPORTED FOR EPIDEMIOLOGIC PURPOSES ONLY.

PCR Result
KPC (bla-KPC) gene DNA Detected

Molecular test

NDM-1 (bla-NDM-1) gene DNA Not Detected
OXA-48 (bla-OXA-48 like) gene DNA Not Detected
VIM (bla-VIM) gene DNA Not Detected
KPC, NDM, OXA-48, and VIM are the most common carbapenemases in the United States, however there are other less common carbapenemases and other mechanisms of carbapenemase resistance not detected by this PCR assay.

IMP PCR Result
IMP (bla-IMP) gene DNA Not Detected

CP-CRE Case Classification

Confirmed CP-CRE

- *Klebsiella spp., E. coli, Enterobacter spp.*
 - Positive phenotypic test or
 - Positive carbapenem resistance mechanism

Suspect CP-CRE

- *Klebsiella spp., E. coli, Enterobacter spp.*
 - Resistance to at least 1 carbapenem
 - No phenotypic or molecular testing done

Not a Case

- BOL report is negative for phenotypic and molecular tests
- All carbapenems are susceptible (MICs don't match case definition)
- Not *Enterobacteriaceae*

Does this Isolate Meet Reporting Requirements?

- ✓ *Klebsiella pneumoniae*
- ✓ Carbapenemase production
- ✓ KPC carbapenemase gene detected
- ✓ Ertapenem MIC ≥ 4
- ✓ Meropenem MIC ≥ 16

= Confirmed CP-CRE Case

$\geq 100,000$ CFU/ml *Klebsiella pneumoniae*, see comment

Comment: Carbapenem resistant Enterobacteriaceae. Carbapenemase producer. KPC detected.

<u>Antibiotic</u>	<u>MIC</u>	<u>Interpretation</u>
Ampicillin	≥ 32	Resistant
Ampicillin/sulbactam	≥ 32	Resistant
Aztreonam	≥ 64	Resistant
Cefazolin	≥ 64	Resistant
Cefepime	2	Resistant
Ceftriaxone	8	Resistant
Ertapenem	≥ 4	Resistant
Gentamicin	≤ 2	Sensitive
Levofloxacin	≤ 1	Sensitive
Meropenem	≥ 16	Resistant
Piperacillin/tazobactam	64	Intermediate
Tobramycin	≤ 2	Sensitive
Trimethoprim/sulfamethoxazole	≤ 2	Sensitive

Does this Isolate Meet Reporting Requirements?

- ✓ *Enterobacter cloacae*
 - X No phenotypic or molecular carbapenemase testing reported
 - ✓ Ertapenem MIC = 2
 - X Meropenem MIC = 1
- = Suspect CP-CRE Case

≥100,000 CFU/ml *Enterobacter cloacae*, see comment

Comment: Carbapenem resistant Enterobacteriaceae.

<u>Antibiotic</u>	<u>MIC</u>	<u>Interpretation</u>
Ampicillin	≥32	Resistant
Ampicillin/sulbactam	≥32	Resistant
Aztreonam	≥64	Resistant
Cefazolin	≥64	Resistant
Cefepime	2	Resistant
Ceftriaxone	8	Resistant
Ertapenem	2	Resistant
Gentamicin	≤2	Susceptible
Levofloxacin	≤1	Susceptible
Meropenem	1	Susceptible
Piperacillin/tazobactam	64	Intermediate
Tobramycin	≤2	Susceptible
Trimethoprim/sulfamethoxazole	≤2	Susceptible

Does this Isolate Meet Reporting Requirements?

✓ *Escherichia coli*

X No phenotypic or molecular carbapenemase testing reported

? Ertapenem and meropenem reported as 'Resistant' but no MIC value reported

= Can not tell if it meets the case definition or not

≥100,000 CFU/ml *Escherichia coli*, see comment

Comment: Carbapenem resistant Enterobacteriaceae.

<u>Antibiotic</u>	<u>MIC</u>	<u>Interpretation</u>
Ampicillin	≥32	Resistant
Ampicillin/sulbactam	≥32	Resistant
Aztreonam		Resistant
Cefazolin		Resistant
Cefepime		Resistant
Ceftriaxone		Resistant
Ertapenem		Resistant
Gentamicin	≤2	Sensitive
Levofloxacin	≤1	Sensitive
Meropenem		Resistant
Piperacillin/tazobactam	64	Intermediate
Tobramycin	≤2	Sensitive
Trimethoprim/sulfamethoxazole	≤2	Sensitive

Tips for CP-CRE Reporting

- **Confirm the organism identification**
 - *Klebsiella* spp., *Escherichia coli*, *Enterobacter* spp.
- **Check for phenotypic carbapenemase testing**
 - ‘Carbapenemase detected’ or ‘Carbapenemase not detected’
 - Confirm the method used: mCIM, CarbaNP, MBL test
- **Check for molecular carbapenemase testing**
 - KPC, NDM, OXA-48, VIM, IMP
- **Confirm carbapenem MIC values**
 - Doripenem, imipenem, or meropenem ≥ 4 $\mu\text{g/ml}$; or ertapenem ≥ 2 $\mu\text{g/ml}$
 - If there are no MIC values reported (e.g., “Resistant”) or no carbapenems reported in MDSS, call the laboratory and ask to speak to a bench technologist

CP-CRE Case Investigation Form

- CRE Investigation Form Sections
 - Patient Information
 - Demographics
 - Laboratory Testing
 - Clinical Information
 - Antimicrobial Therapy

<input type="button" value="Submit Changes"/> <input type="button" value="Cancel Changes"/> <input type="button" value="Print"/>					
<h2>CRE Investigation Form</h2> <p>Carbapenemase-Producing Carbapenem-Resistant Enterobacteriaceae (CP-CRE) Michigan Department of Health and Human Services Communicable Disease Division</p>					
Investigation Information					
Investigation ID	Onset Date <i>mm/dd/yyyy</i>	Diagnosis Date <i>mm/dd/yyyy</i>	Referral Date <i>mm/dd/yyyy</i>	Case Entry Date <i>mm/dd/yyyy</i>	Case Completion Date <i>mm/dd/yyyy</i>
	<input type="text"/>	<input type="text"/>	<input type="text"/>	<input type="text" value="04/05/2018"/>	<input type="text"/>
Investigation Status	Case Status				<input type="checkbox"/> State Prison Case
New	<input type="radio"/> Confirmed <input type="radio"/> Not a Case <input type="radio"/> Probable <input type="radio"/> Suspect <input type="radio"/> Unknown <input type="radio"/> Non-Michigan Case				
Patient Status	Patient Status Date <i>mm/dd/yyyy</i>	Case Disposition	Part of an outbreak?	Outbreak Name	Case Updated Date <i>mm/dd/yyyy</i>
	<input type="text" value="04/05/2018"/>	<input type="text"/>	<input type="text"/>	<input type="text"/>	<input type="text" value="04/05/2018"/>
Facility where specimen collected:			Participate in the CRE Surveillance and Prevention Initiative		
<input type="text"/>			<input type="radio"/> Yes <input type="radio"/> No		
Patient Information					
Patient ID	First	Last	Middle		
<input type="text"/>	<input type="text"/>	<input type="text"/>	<input type="text"/>		
Street Address					
<input type="text"/>					
City	County	State	Zip		
<input type="text"/>	<input type="text"/>	<input type="text"/>	<input type="text"/>		
Home Phone ### ### ####	Ext.	Other Phone ### ### ####	Ext.		
<input type="text"/>	<input type="text"/>	<input type="text"/>	<input type="text"/>		
Parent/Guardian (required if under 18)					
First	Last	Middle			
<input type="text"/>	<input type="text"/>	<input type="text"/>			
Demographics					

CP-CRE Laboratory Testing

- **Laboratory Testing information is required to determine case classification**

- Date collected
- Specimen source
- Organism
- MIC
 - need actual numerical value
- Carbapenemase testing
 - e.g., mCIM, CarbaNP
- Resistance mechanism testing
 - e.g. PCR, Carba-R

Laboratory Testing and Microbiology Information		
Date Specimen Collected (mm/dd/yyyy)	Specimen Type: <input type="radio"/> Clinical Culture <input type="radio"/> Surveillance Culture	
Specimen Source: <input type="checkbox"/> Blood <input type="checkbox"/> Respiratory/Spitum <input type="checkbox"/> Urine <input type="checkbox"/> Wound, skin, or soft tissue <input type="checkbox"/> Rectal or perianal <input type="checkbox"/> Other, specify _____		
Organism: <input type="checkbox"/> <i>Klebsiella</i> species, specify _____ <input type="checkbox"/> <i>Escherichia coli</i> <input type="checkbox"/> <i>Enterobacter</i> species, specify _____ <input type="checkbox"/> Other, specify _____		
Date Antimicrobial Testing Results Reported (mm/dd/yyyy)		
Antimicrobial Susceptibility Testing Results (fill only those that were reported):		
Antimicrobial	Minimum Inhibitory Concentration (MIC) (mg/L)	Interpretation (S, susceptible; I, intermediate; R, resistant)
Imipenem		
Meropenem		
Doripenem		
Ertapenem		
Amikacin		
Tobramycin		
Gentamicin		
Tigecycline		
Colistin		
Phenotypic Carbapenemase Production Testing		
Test Method:	If other, specify:	Result: <input type="radio"/> Positive <input type="radio"/> Negative <input type="radio"/> Indeterminate
Resistance Mechanism for Carbapenemase Testing: (e.g., PCR or other molecular genetic test)		
Resistance Mechanism	Response	
KPC	<input type="radio"/> Detected <input type="radio"/> Not Detected <input type="radio"/> Not Tested <input type="radio"/> Invalid	
NDM	<input type="radio"/> Detected <input type="radio"/> Not Detected <input type="radio"/> Not Tested <input type="radio"/> Invalid	
OXA-48	<input type="radio"/> Detected <input type="radio"/> Not Detected <input type="radio"/> Not Tested <input type="radio"/> Invalid	
VIM	<input type="radio"/> Detected <input type="radio"/> Not Detected <input type="radio"/> Not Tested <input type="radio"/> Invalid	
IMP	<input type="radio"/> Detected <input type="radio"/> Not Detected <input type="radio"/> Not Tested <input type="radio"/> Invalid	
Other, specify _____	<input type="radio"/> Detected <input type="radio"/> Not Detected <input type="radio"/> Not Tested <input type="radio"/> Invalid	

CP-CRE Clinical Info

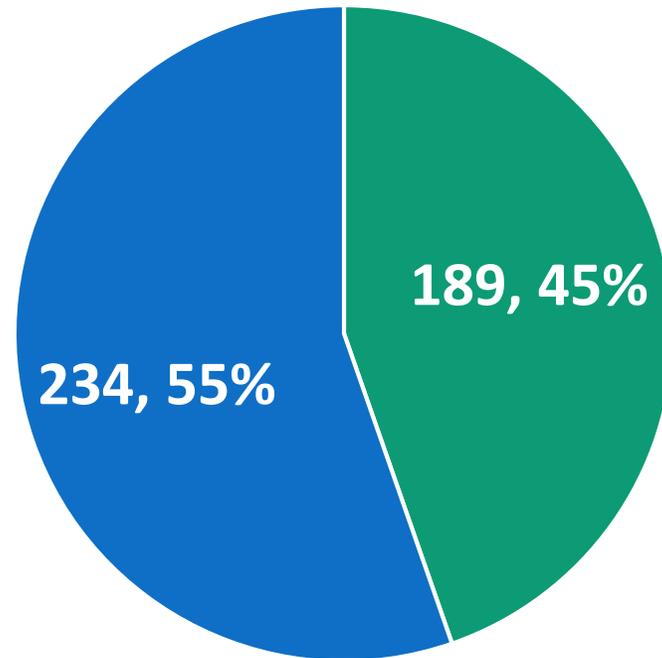
- Healthcare exposures
 - Acute care, long-term care
- Travel
 - Location
 - Healthcare abroad

Clinical Information	
Date of Patient Admission or Presentation (mm/dd/yyyy) <input type="text"/>	Date Patient was placed in Contact Precautions/Isolation (if an inpatient) (mm/dd/yyyy) <input type="text"/>
Patient Admitted/Presented From: <input type="radio"/> Long-Term Care/Skilled Nursing Facility <input type="radio"/> Outside Acute Care Hospital <input type="radio"/> Home <input type="radio"/> Long-Term Acute Care Hospital <input type="radio"/> Unknown <input type="radio"/> Other, specify <input type="text"/>	
Date of Patient Discharge (mm/dd/yyyy) <input type="text"/>	Was information on CRE status shared with transferring agency and admitting facility: <input type="radio"/> Yes <input type="radio"/> No <input type="radio"/> Unknown
Patient Discharged to: <input type="radio"/> Long-Term Care/Skilled Nursing Facility <input type="radio"/> Outside Acute Care Hospital <input type="radio"/> Home <input type="radio"/> Long-Term Acute Care Hospital <input type="radio"/> Unknown <input type="radio"/> Other, specify <input type="text"/>	
Has Patient previously been hospitalized in an Acute Care Hospital in the last 90 days: <input type="radio"/> Yes <input type="radio"/> No <input type="radio"/> Unknown If Yes, please indicate the facility name and dates of stay (if known) Facility: <input type="text"/> Dates: (mm/dd/yyyy) From <input type="text"/> To <input type="text"/>	
Has Patient been admitted to a Long-Term Acute Care Hospital in the last 90 days: <input type="radio"/> Yes <input type="radio"/> No <input type="radio"/> Unknown If Yes, please indicate the facility name and dates of stay (if known) Facility: <input type="text"/> Dates: (mm/dd/yyyy) From <input type="text"/> To <input type="text"/>	
Has Patient been admitted to a Long-Term Care Facility (e.g., nursing home, SNF) in the last 90 days: <input type="radio"/> Yes <input type="radio"/> No <input type="radio"/> Unknown If Yes, please indicate the facility name and dates of stay (if known) Facility: <input type="text"/> Dates: (mm/dd/yyyy) From <input type="text"/> To <input type="text"/>	
Indwelling Devices (in place within 2 calendar days of specimen collection):	
Central Venous Line: <input type="radio"/> Yes <input type="radio"/> No <input type="radio"/> Unknown	Mechanical Ventilation: <input type="radio"/> Yes <input type="radio"/> No <input type="radio"/> Unknown
<input type="radio"/> Unknown	Wound VAC (vacuum-assisted closure): <input type="radio"/> Yes <input type="radio"/> No <input type="radio"/> Unknown

Does the Patient have a history of any International Travel in the last 6 months? <input type="radio"/> Yes <input type="radio"/> No <input type="radio"/> Unknown If Yes, please indicate the country(ies) and dates of travel Country: <input type="text"/>	
Dates: (mm/dd/yyyy) From <input type="text"/> To <input type="text"/>	
If Yes, please indicate if any healthcare exposures internationally Exposure: <input type="text"/>	

CP-CRE Cases Reported to MDSS

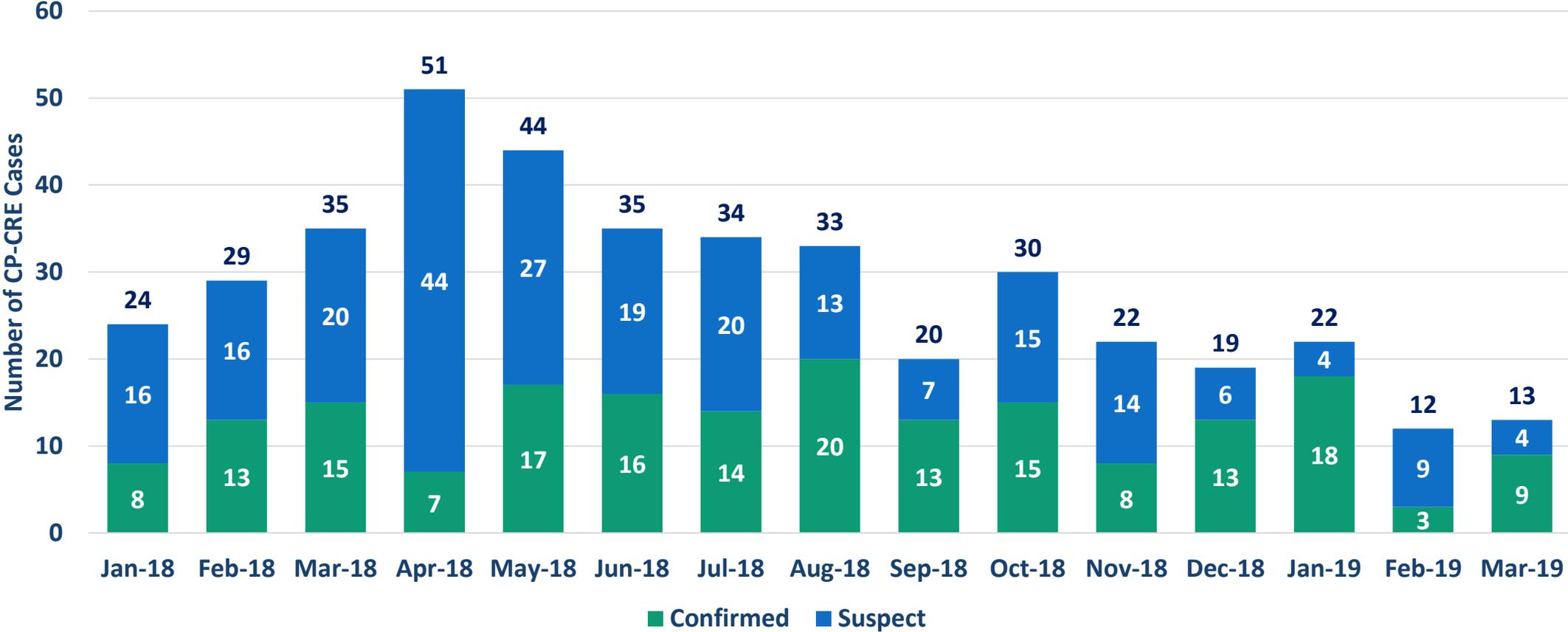
Jan 2018 – Mar 2019



■ Confirmed ■ Suspect

CP-CRE Cases Reported to MDSS

Jan 2018 – Mar 2019



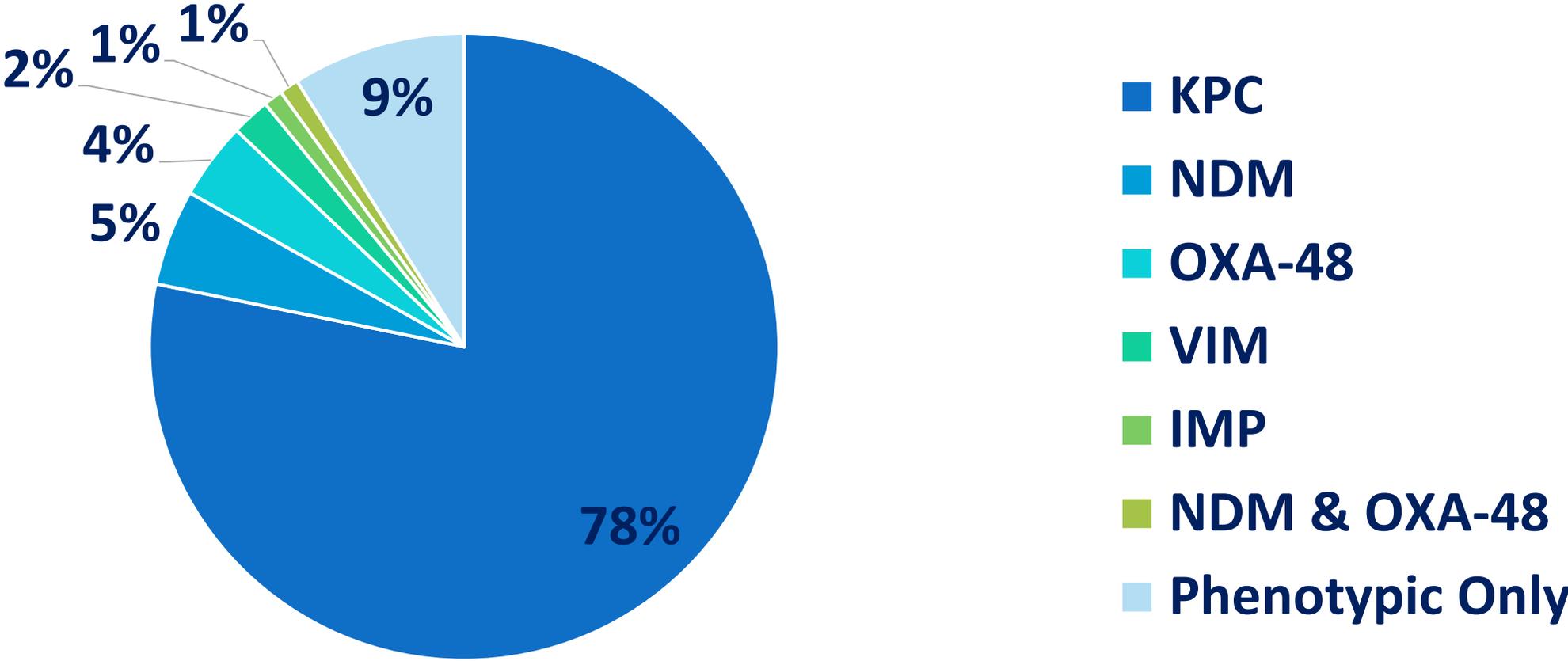
CP-CRE Cases by Organism

Jan 2018 – Mar 2019

Organism	CP-CRE Cases		
	Confirmed (n=189)	Suspect (n=234)	Total (n=423)
<i>Klebsiella spp.</i>	126 (67%)	95 (41%)	221 (52%)
<i>Klebsiella pneumoniae</i>	118	73	191
<i>Klebsiella aerogenes</i>	4	15	19
<i>Klebsiella oxytoca</i>	3	7	10
<i>Klebsiella variicola</i>	1	0	1
<i>Enterobacter spp.</i>	36 (19%)	68 (29%)	104 (25%)
<i>Enterobacter cloacae</i>	36	66	102
<i>Enterobacter asburiae</i>	0	1	1
<i>Enterobacter hormaechei</i>	0	1	1
<i>Escherichia coli</i>	27 (14%)	71 (30%)	98 (23%)

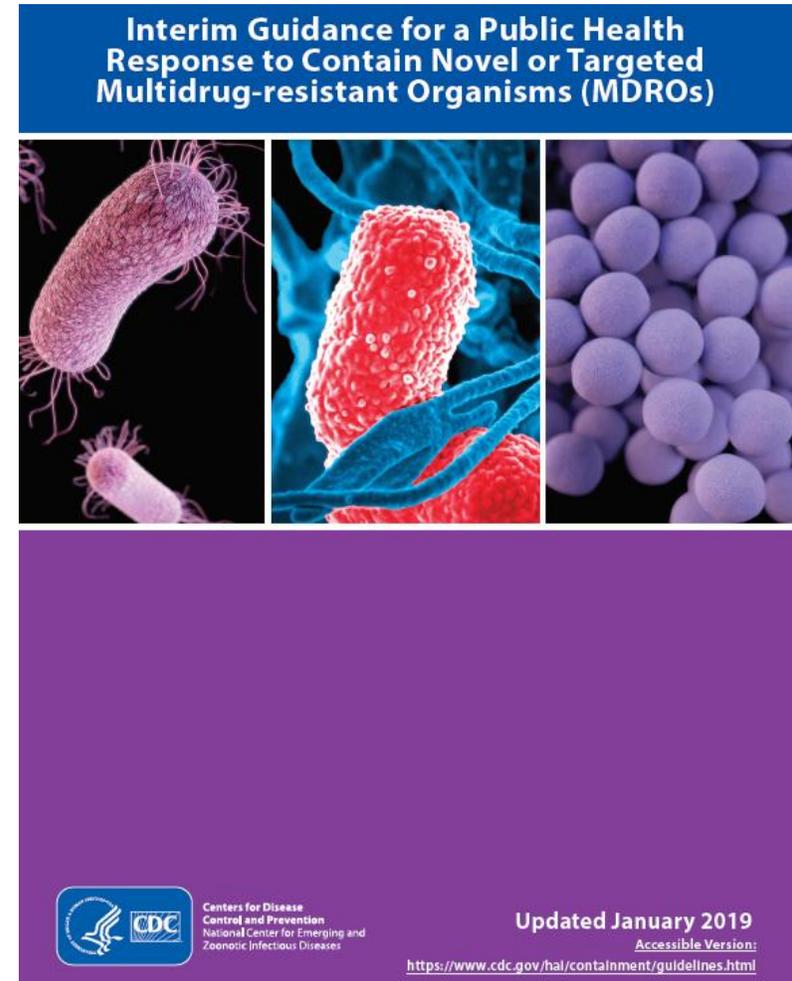
Confirmed CP-CRE Cases by Mechanism

Jan 2018 – Mar 2019



CDC Guidance for Novel MDRO Containment

- Response to a **single case** of resistance
- Goal to slow the spread of novel or unusual MDROs or resistance mechanisms
- Healthcare Contact Investigation
 - Notification/communication of status
 - Prompt implementation of precautions
 - Screening roommates
 - Screening broader healthcare contacts
 - Prospective and retrospective lab surveillance
- Assessment of infection control practices



<https://www.cdc.gov/hai/containment/guidelines.html>

Considerations for CP-CRE Screening

- **Who to screen**

- Epi-linked patient contacts of newly identified CP-CRE patients
- Patients with an overnight stay in a healthcare facility outside the U.S. in the prior 6-12 months
- Patients at high-risk for colonization

- **Colonization screen testing**

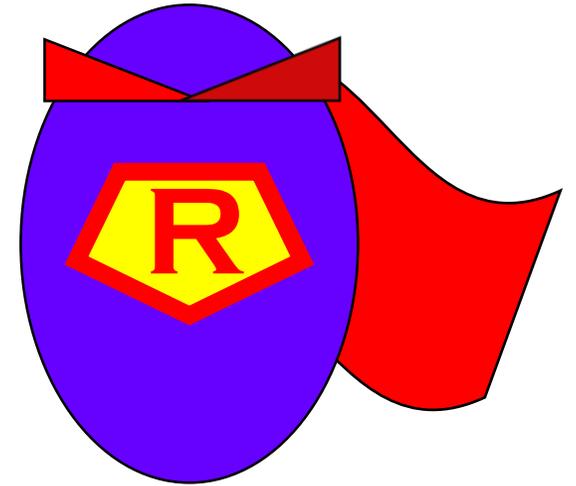
- Available through MDHHS BOL
- Rectal swab collection
- Collection supplies are available for facilities (e.g., SNF)

Candida auris



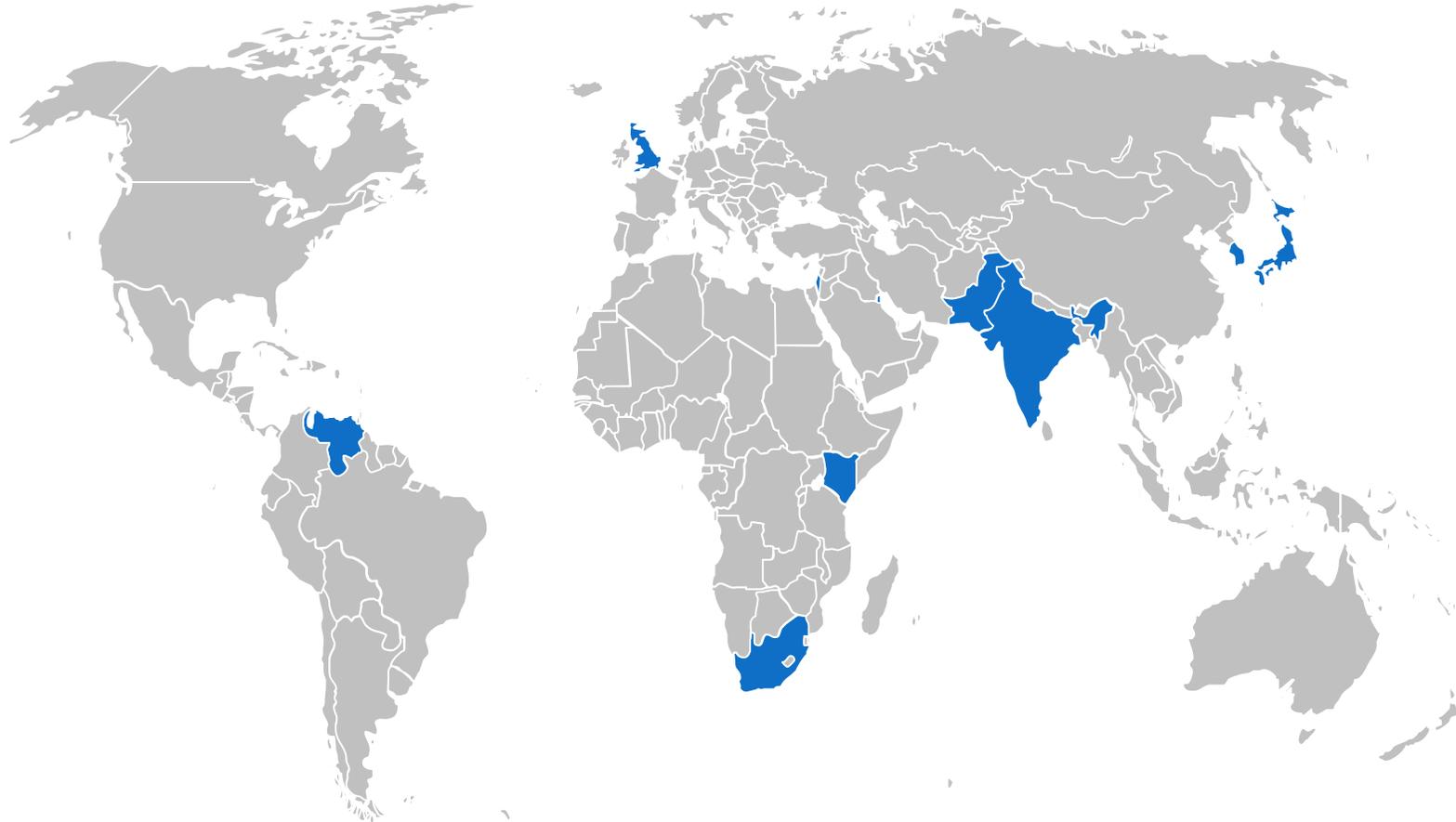
Candida auris is a Public Health Concern

- Recent worldwide emergence
- Highly drug-resistant yeast
- It can cause serious, invasive infections associated with high-mortality rates
- It is difficult to identify
- It can spread easily in healthcare settings



A Fungal Superbug!

Discovery of *Candida auris*



2009

Japan

2010

2011

**South
Korea**

2012

2013

India

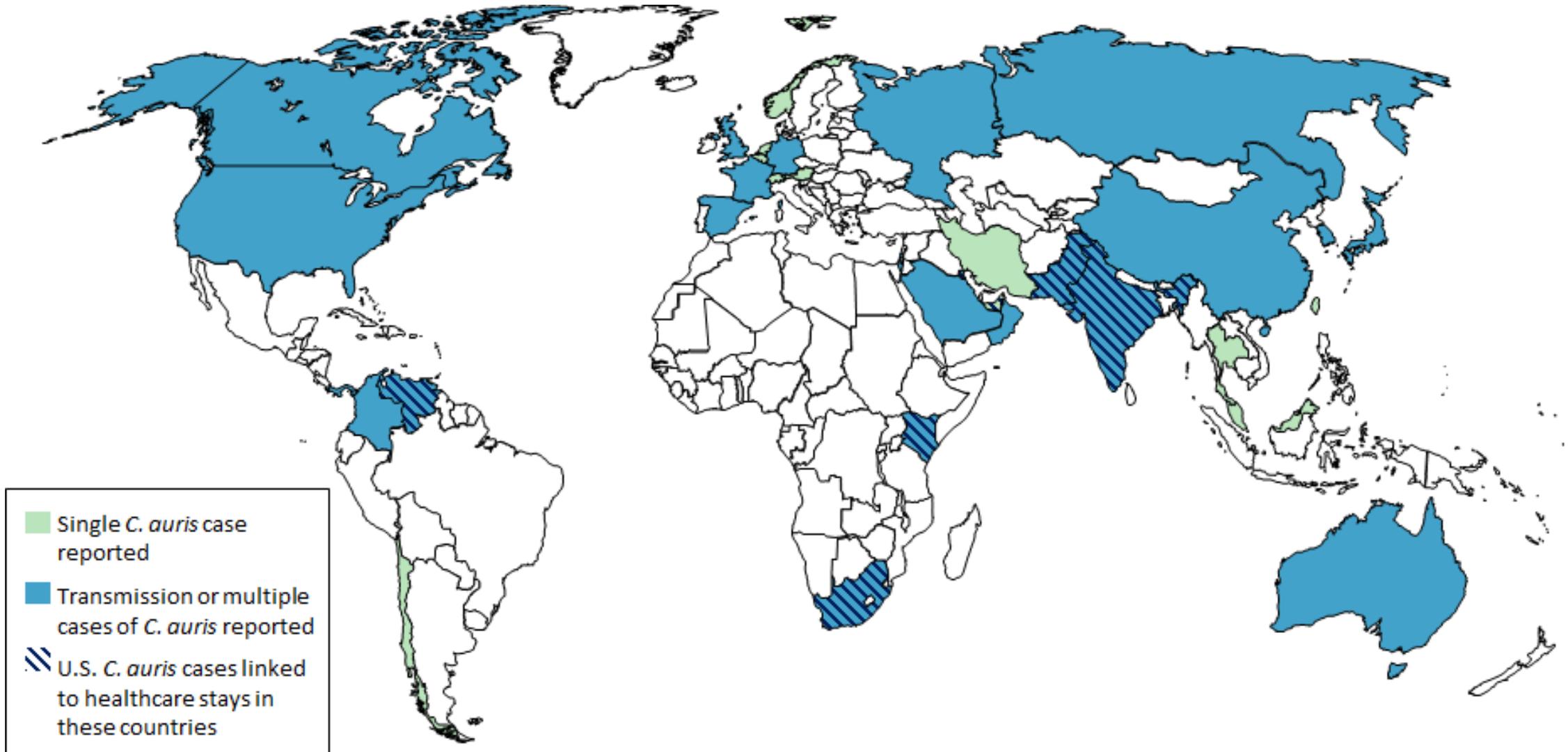
2014

**South Africa
Kenya
Kuwait**

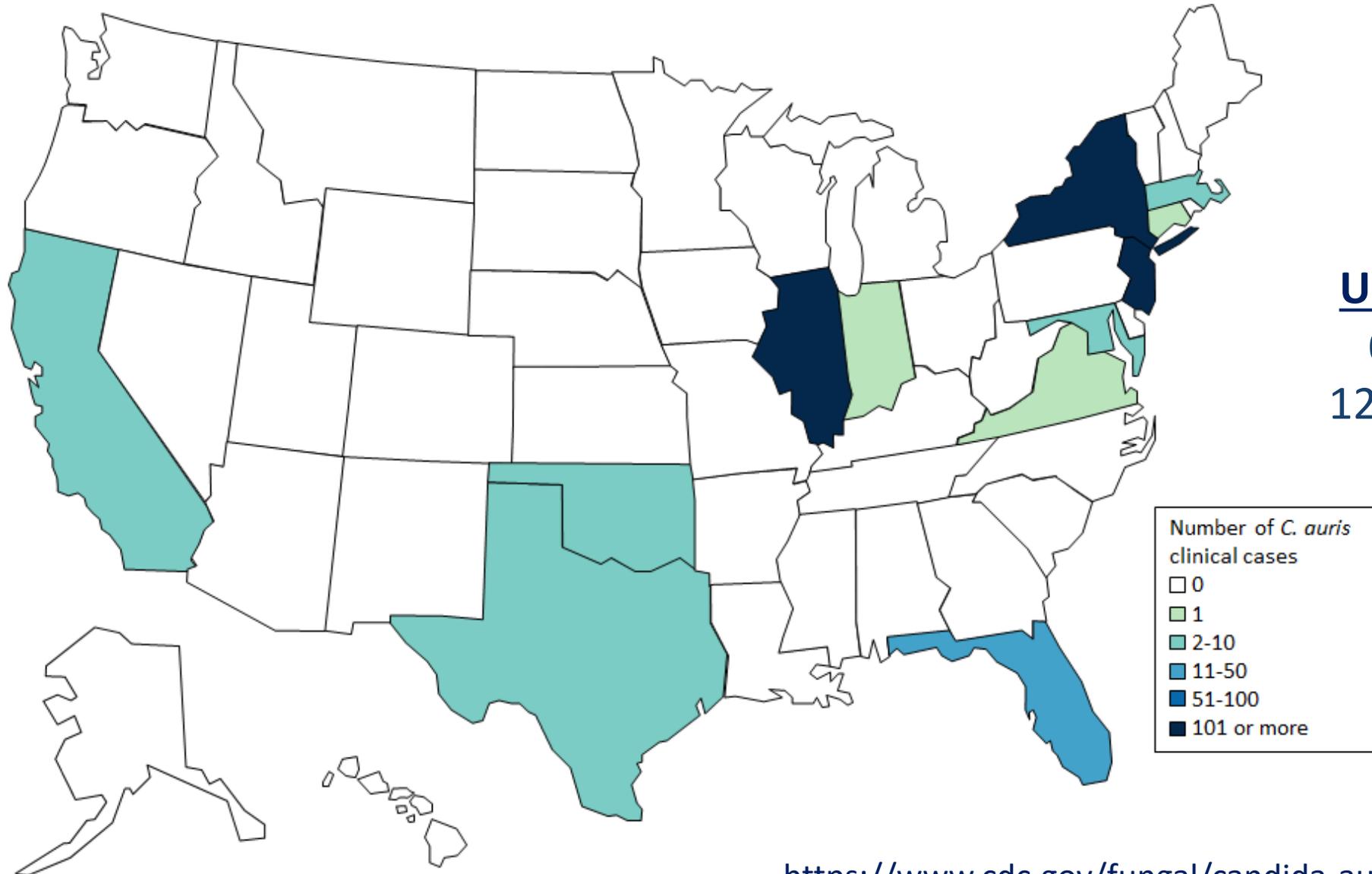
2015

**Pakistan
Venezuela
Israel
UK**

Reports of *Candida auris*, April 2019



Clinical Cases of *Candida auris*, April 2019



U.S. Cases Reported
654 Clinical Cases
1207 Screening Cases

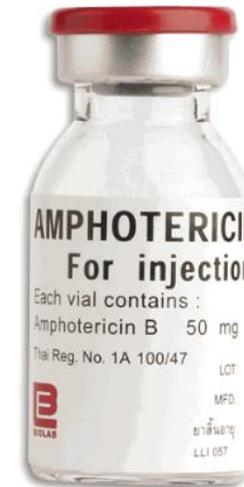
Antifungal Resistance is Common



>90%
Azoles



7%
Echinocandins

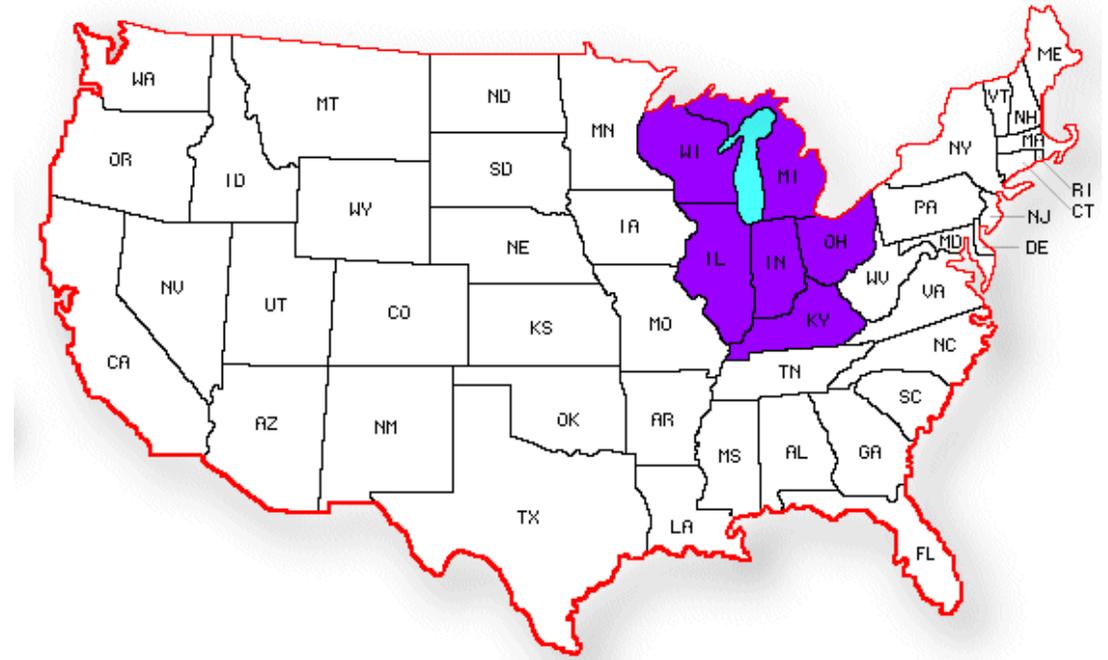


35%
Polyenes

- >40% multi-drug resistant
- A few isolates resistant to all three classes

Midwest Antifungal Resistance

- 2019 Q1 testing of 78 isolates:
 - 23.1% resistant to fluconazole
 - 0% resistant to amphotericin B
 - 2.6% resistant to echinocandins



Candida auris Causes Invasive Infections

- ~50% of clinical cases had bloodstream infections
- Candidemia associated with 30-60% mortality rates



Risk Factors for *Candida auris*



Older age



**Multiple
healthcare
exposures**

Long-term care

Acute care

Outside US



**Antifungals
and
antibiotics**



**Indwelling
devices**

Catheters

Ventilator

PEG tube



**Comorbid
conditions**

Cancer

Diabetes

Renal Disease

Recent surgery

Candida auris Identification Can Be Difficult

- Phenotypic characteristics are not sufficient for identification
- Many yeast identification methods may not correctly identify *C. auris*
 - Misidentification of species
 - No identification
- Yeast isolates may not be identified, especially in non-invasive sites
 - >40% of U.S. clinical cases were identified from non-bloodstream isolates (e.g., urine, wounds)



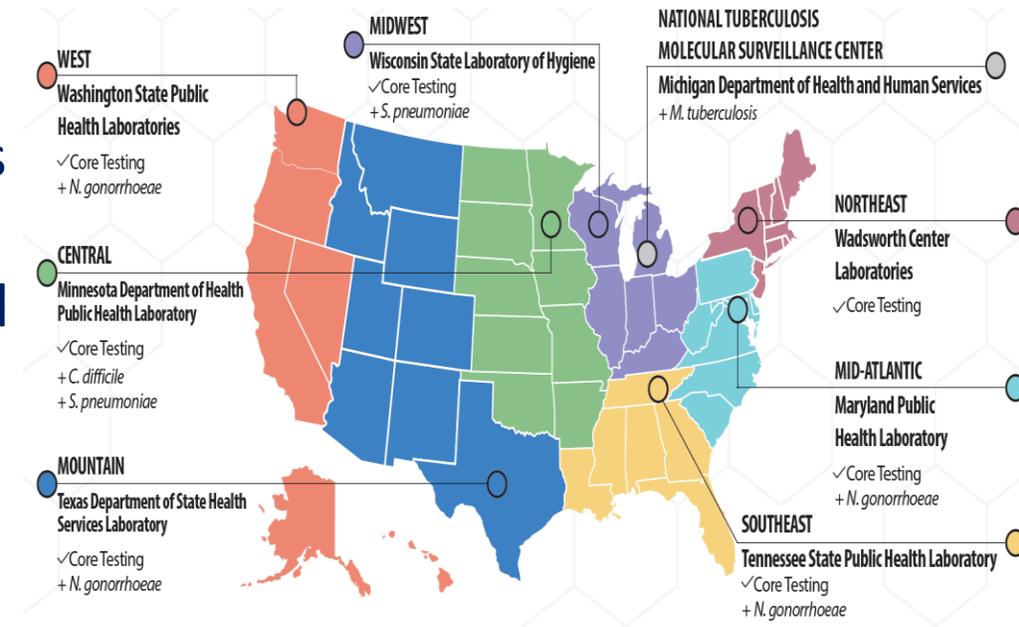
Candida auris on CHROMagar Candida, displaying multiple color morphs

Identification Method	Database/Software, if applicable	<i>C. auris</i> is confirmed if initial identification is <i>C. auris</i> .	<i>C. auris</i> is possible if the following initial identifications are given. Further work-up is needed to determine if the isolate is <i>C. auris</i> .
Bruker Biotyper MALDI-TOF	RUO libraries (Versions 2014 [5627] and more recent)	<i>C. auris</i>	n/a
	CA System library (Version Claim 4)	<i>C. auris</i>	n/a
bioMérieux VITEK MS MALDI-TOF	RUO library (with Saramis Version 4.14 database and Saccharomycetaceae update)	<i>C. auris</i>	<i>C. haemulonii</i> No identification
	IVD library	n/a	<i>C. haemulonii</i> No identification
VITEK 2 YST	Software version 8.01	<i>C. auris</i>	<i>C. haemulonii</i> <i>C. duobushaemulonii</i> <i>Candida</i> spp. not identified
	Older versions	n/a	<i>C. haemulonii</i> <i>C. duobushaemulonii</i> <i>Candida</i> spp. not identified
API 20C		n/a	<i>Rhodotorula glutinis</i> (with characteristic red color present) <i>C. sake</i> <i>Candida</i> spp. not identified
BD Phoenix		n/a	<i>C. catenulata</i> <i>C. haemulonii</i> <i>Candida</i> spp. not identified
MicroScan		n/a	<i>C. lusitaniae</i> * <i>C. guilliermondii</i> * <i>C. parapsilosis</i> * <i>C. famata</i> <i>Candida</i> spp. not identified
RapID Yeast Plus		n/a	<i>C. parapsilosis</i> * <i>Candida</i> spp. not identified

* *C. guilliermondii*, *C. lusitaniae*, and *C. parapsilosis* generally make hyphae or pseudohyphae on cornmeal agar. If hyphae or pseudohyphae are not present on cornmeal agar, the isolate should raise suspicions of being *C. auris* as *C. auris* typically does not make hyphae or pseudohyphae. However, some *C. auris* isolates have formed hyphae or pseudohyphae. Therefore, it would be prudent to consider any *C. guilliermondii*, *C. lusitaniae*, and *C. parapsilosis* isolates identified on MicroScan and any *C. parapsilosis* isolates identified on RapID Yeast Plus as possible *C. auris* isolates and further work-up should be considered.

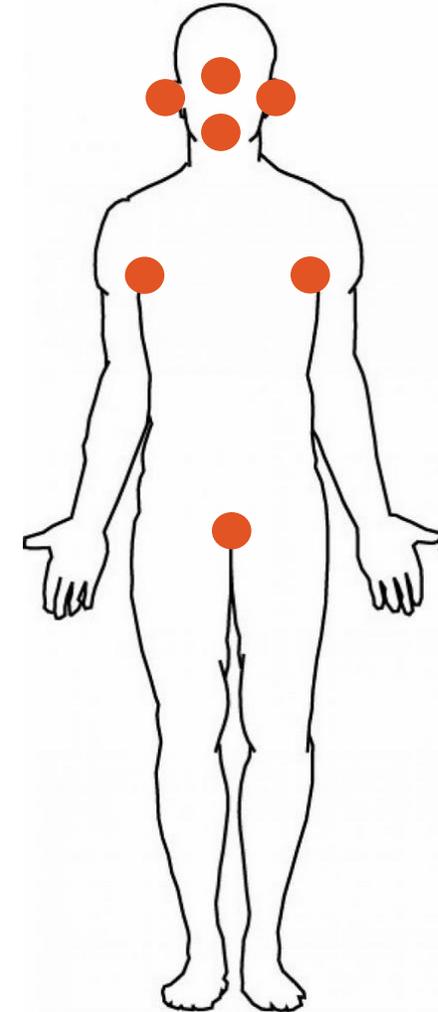
Candida Testing Available

- Labs can send isolates to MDHHS BOL → ARLN
- Isolates to send:
 - Any confirmed or suspected *Candida auris* isolates
 - Any *Candida* species that was unable to be identified after a validated method was attempted
 - *Candida* from normally sterile sites (except *C. albicans* or *C. glabrata*)
 - Multi-drug resistant *Candida* isolates
 - Unusual *Candida* species



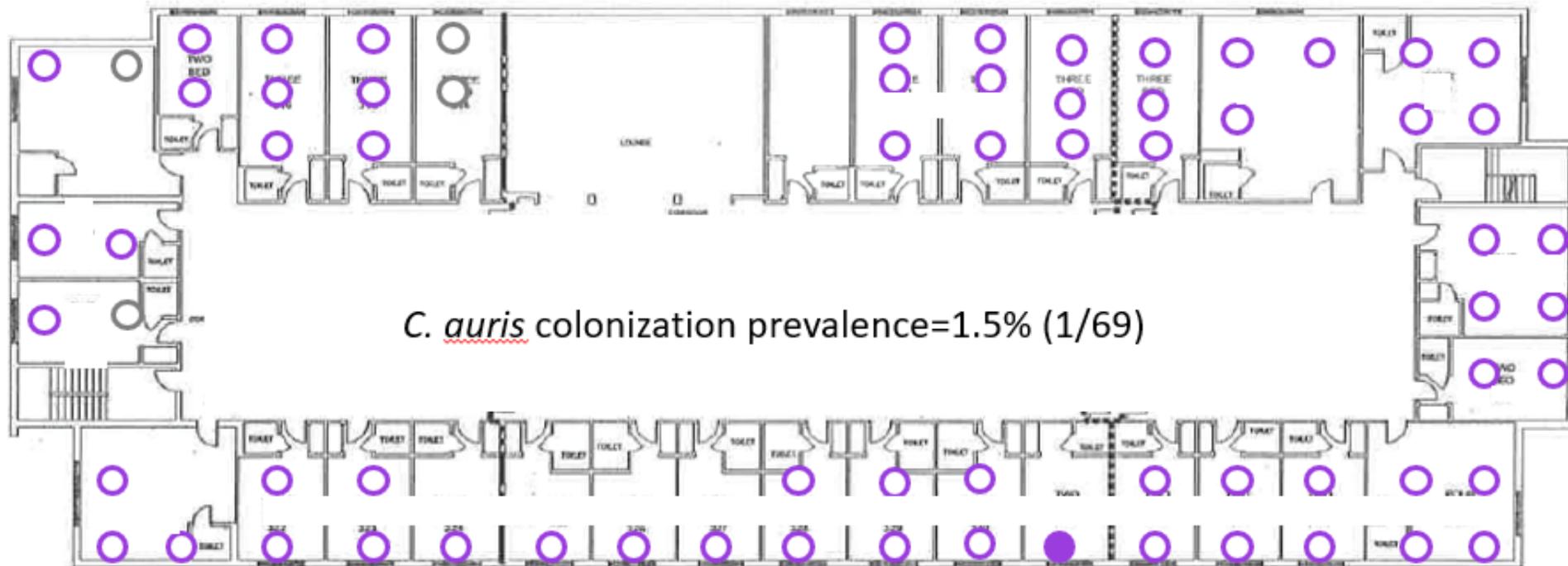
Candida auris Colonizes Skin & Other Body Sites

- **Colonization is a risk for:**
 - Invasive infections
 - Transmission to other patients
- **Duration of colonization can be prolonged**
- **Higher rates of colonization in high-acuity long-term care facilities**



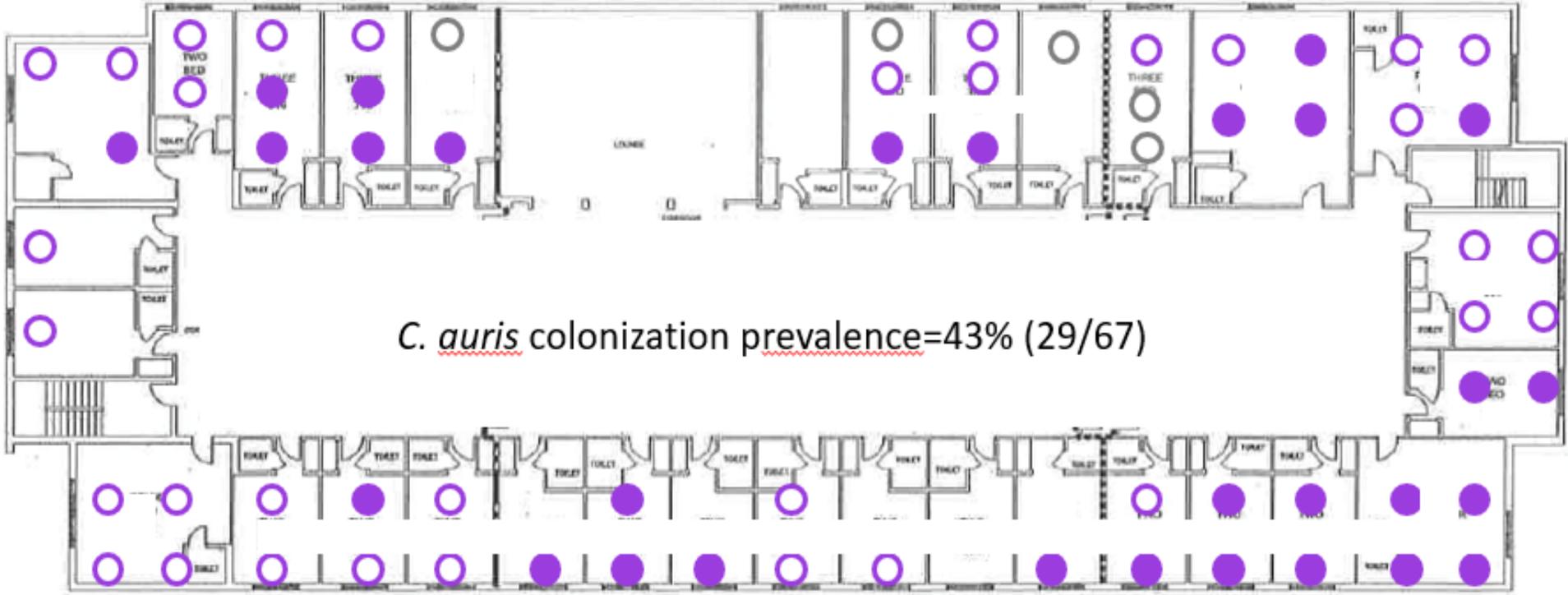
vSNF A Ventilator/Trach Floor

March 2017 *C. auris* PPS Results



- *C. auris* positive
- Screened negative for *C. auris*
- Not tested for *C. auris* (refused or not in room)

vSNF A Ventilator/Trach Floor January 2018 *C. auris* PPS Results

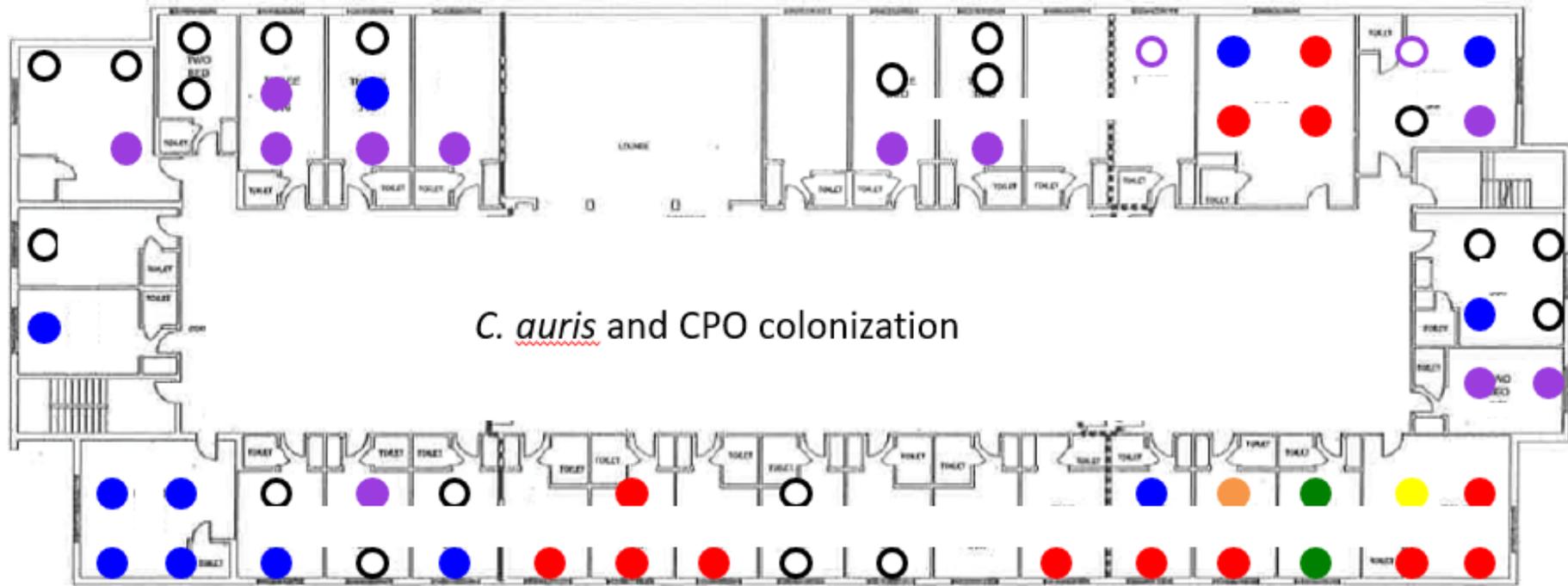


C. auris colonization prevalence=43% (29/67)

- *C. auris* positive
- Screened negative for *C. auris*
- Not tested for *C. auris* (refused or not in room)

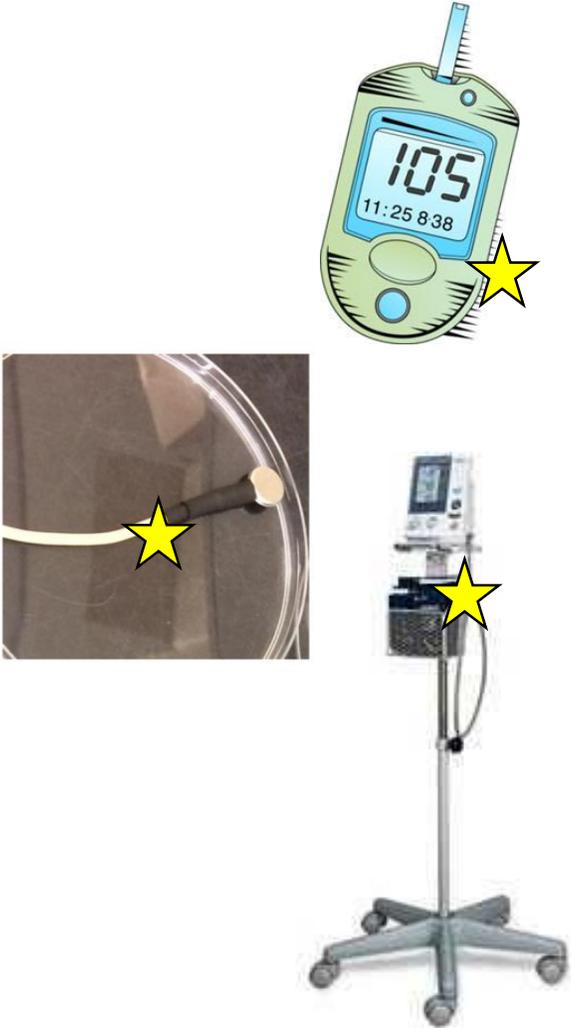
Slide courtesy of Chicago Department of Public Health.

vSNF A Ventilator/Trach Floor January 2018 CPO and *C. auris* PPS Results

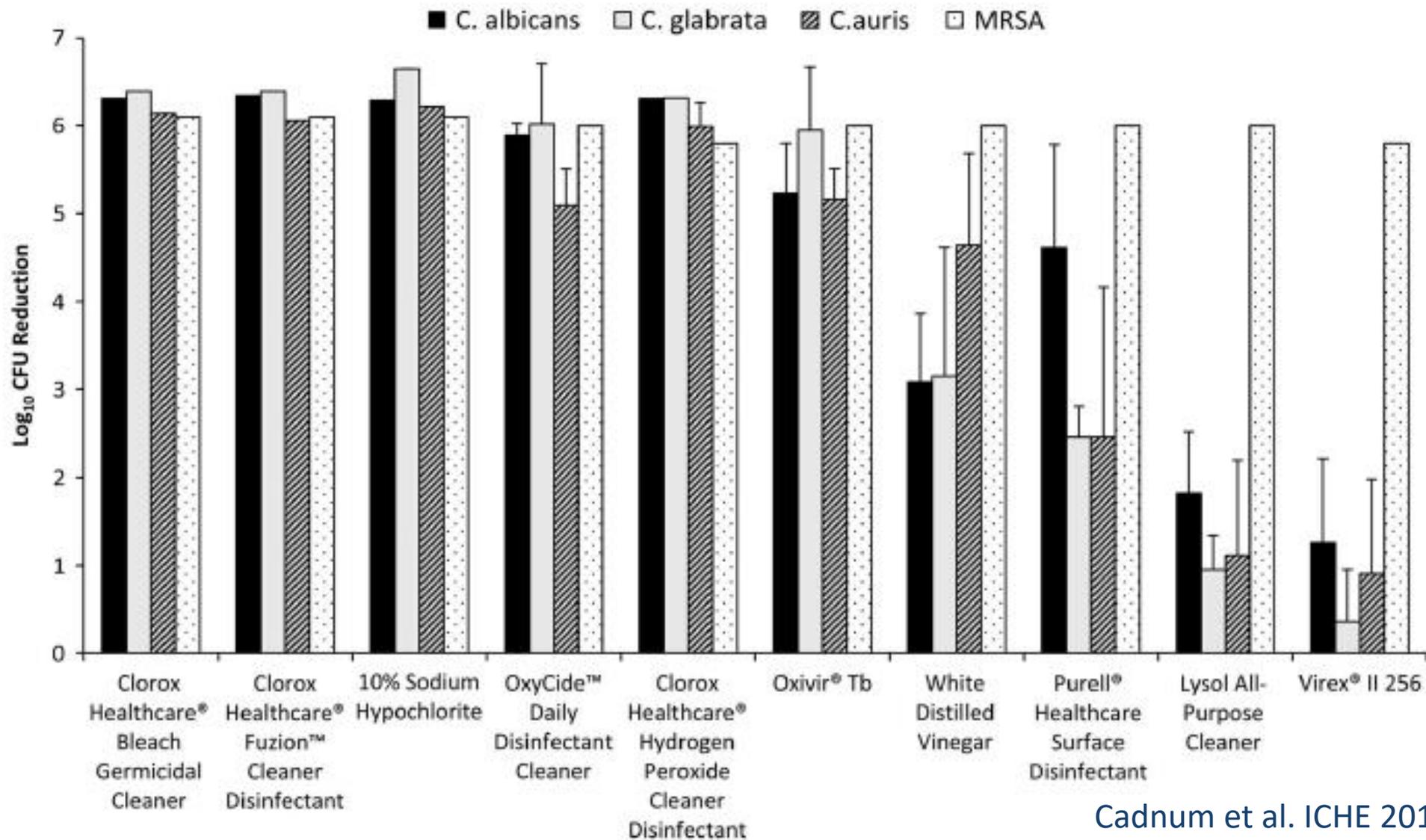


- *C. auris*
- *C. auris* and KPC
- KPC or CRE with unknown mechanism of resistance
- *C. auris*, KPC, and NDM
- *C. auris*, VIM-CRPA, and KPC
- *C. auris* and KPC-CRPA
- Screened negative for *C. auris*, but not tested for CRE
- Screened negative for CRE and *C. auris*

Candida auris Can Persist in the Environment



Routine Disinfectants Not as Effective



Candida auris is Now a Reportable Condition

- June 2018 - CSTE position statement for standardized *C. auris* case definition and national notification of cases was passed
- January 1, 2019 - *C. auris* was made a reportable condition in Michigan

Candida auris Reporting Requirements

- Please report any laboratory finding that meets either of the following criteria:
 - Detection of *C. auris* in a specimen using either culture or a culture-independent diagnostic test (CIDT) (e.g., Polymerase Chain Reaction [PCR])
 - Detection of an organism that commonly represents a *C. auris* misidentification in a specimen by culture (i.e., *Candida haemulonii*)
- Laboratories shall immediately submit **confirmed or suspect *C. auris*** isolates, subcultures, or specimens from the patient being tested to the MDHHS Lansing laboratory (**Required**)

Candida auris Reporting in MDSS

- **At this time, base case form is available**
 - Expecting to have a *C. auris* specific form by late 2019/early 2020
- **Important information to collect:**
 - Demographics
 - Laboratory testing results
 - Where specimen was collected
 - Healthcare facility exposures
 - Long-term care residence
 - Tracheostomy/ventilator use
 - Travel history: International and Domestic
 - MDRO status

<input type="button" value="Submit Changes"/> <input type="button" value="Cancel Changes"/> <input type="button" value="Print"/>					
<h2>Base Case Investigation Report</h2> <p>Michigan Department of Health and Human Services Communicable Disease Division</p>					
Investigation Information					
Reportable Condition					
<input type="radio"/> Anthrax		<input type="radio"/> Botulism, Other		<input type="radio"/> Chancroid	
<input type="radio"/> Granuloma Inguinale		<input type="radio"/> Head Lice		<input type="radio"/> Hemorrhagic Fever	
<input type="radio"/> Lymphogranuloma venereum		<input type="radio"/> Melioidosis		<input type="radio"/> Rabies, Human	
<input type="radio"/> Shingles		<input type="radio"/> Staphylococcus Aureus Infection		<input type="radio"/> Strep Throat	
<input type="radio"/> Trachoma		<input type="radio"/> Typhus		<input type="radio"/> VZ Infection, Unspecified	
<input type="radio"/> Unusual Outbreak or Occurrence: <input type="text"/>		<input type="radio"/> Candida auris			
<input type="radio"/> Novel Coronavirus					
Investigation ID	Onset Date <small>mm/dd/yyyy</small>	Diagnosis Date <small>mm/dd/yyyy</small>	Referral Date <small>mm/dd/yyyy</small>	Case Entry Date <small>mm/dd/yyyy</small>	Case Completion Date <small>mm/dd/yyyy</small>
<input type="text"/>	<input type="text"/>	<input type="text"/>	<input type="text"/>	04/04/2019	<input type="text"/>
Investigation Status		Case Status			
New		<input type="radio"/> Confirmed <input type="radio"/> Not a Case <input type="radio"/> Probable <input type="radio"/> Suspect <input type="radio"/> Unknown <input type="radio"/> Non-Michigan Case <input type="checkbox"/> State Prison Case			
Patient Status	Patient Status Date <small>mm/dd/yyyy</small>	Case Disposition	Part of an outbreak?	Outbreak Name	Case Updated Date <small>mm/dd/yyyy</small>
Alive	04/04/2019	<input type="text"/>	<input type="text"/>	<input type="text"/>	04/04/2019
Patient Information					
Patient ID	First	Last	Middle		
<input type="text"/>	<input type="text"/>	<input type="text"/>	<input type="text"/>		
Street Address					
<input type="text"/>					

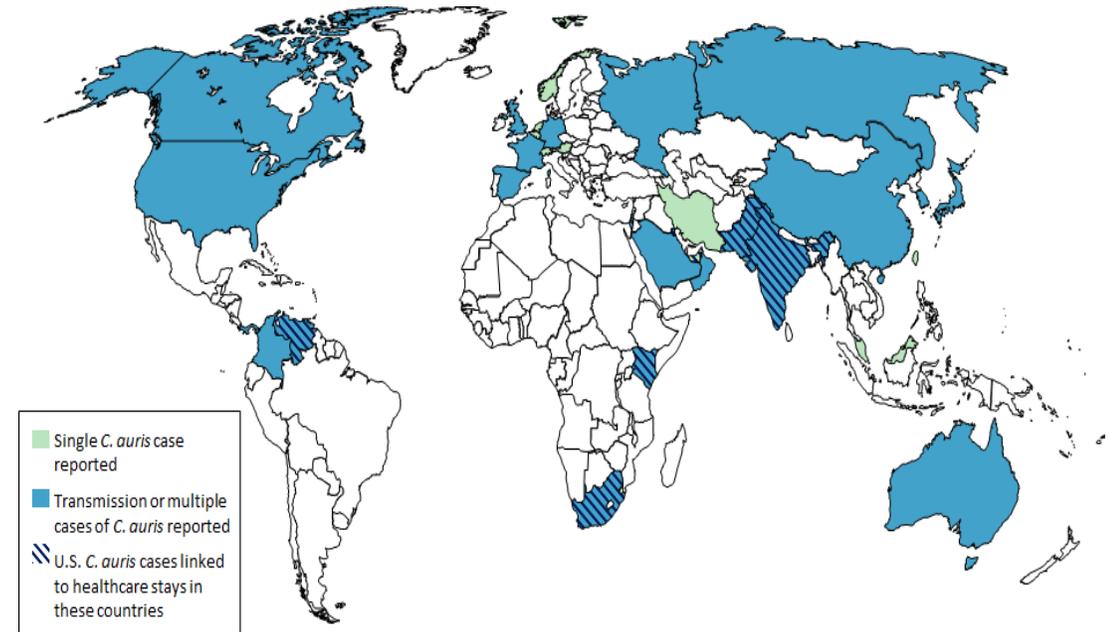
CDC Guidance for Novel MDRO Containment

- Response to a **single case** of resistance
- Goal to **slow the spread** of novel or unusual MDROs or resistance mechanisms
- **Contact Investigation**
 - Screening roommates
 - Screening broader healthcare contacts
 - Prospective and retrospective lab surveillance
 - Assessment of infection control practices



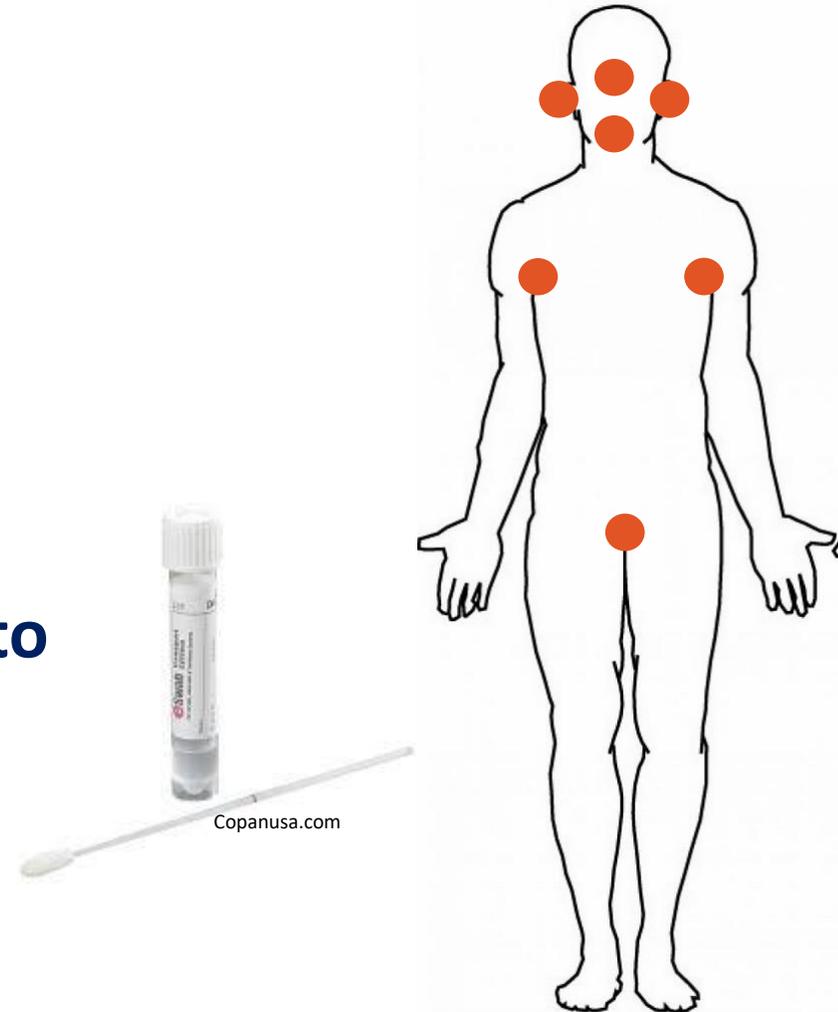
Screening Recommendations

- **Close contacts of *C. auris* patients**
- **Patients with overnight stay in a healthcare facility outside the U.S. in past 12 months**
- **Patients in high-acuity long-term care facilities (e.g., care for ventilated patients), especially those with CP-CRE or other MDROs**



Candida auris Colonization Testing

- **Must be approved by SHARP unit**
 - We can assist with obtaining specimen collection kits, contact investigation, and follow-up
- **Testing done through ARLN**
- **Axilla/groin swab most sensitive sites to detect colonization**



Infection Prevention for *Candida auris*

- **Place patients in a single-patient room, using Standard and Contact Precautions**
- **Emphasize adherence to hand hygiene**
- **Clean and disinfect patient care environment and reusable equipment**
 - Daily and terminal cleaning with EPA List K products active against *C. difficile*
- **Inter-facility communication at transfer to another healthcare facility**
- **Conduct surveillance to detect new cases and ongoing transmission**

How Healthcare Facilities Can Prepare for *C. auris*

- **Work with your laboratory to ensure your yeast identification method can identify *C. auris***
 - If it cannot, know when to suspect *C. auris* and send suspected isolates to the Bureau of Laboratories for testing through ARLN
- **Establish a surveillance protocol with your lab**
 - Prompt notification and reporting of **confirmed or suspect *C. auris*** isolates to infection prevention and public health
- **Know which patients in your facility are at higher risk for *C. auris***
 - Request colonization screening for high-risk patients through ARLN
- **Develop a facility response plan**
 - Prompt implementation of infection prevention and control measures

How Local Health Departments Can Respond

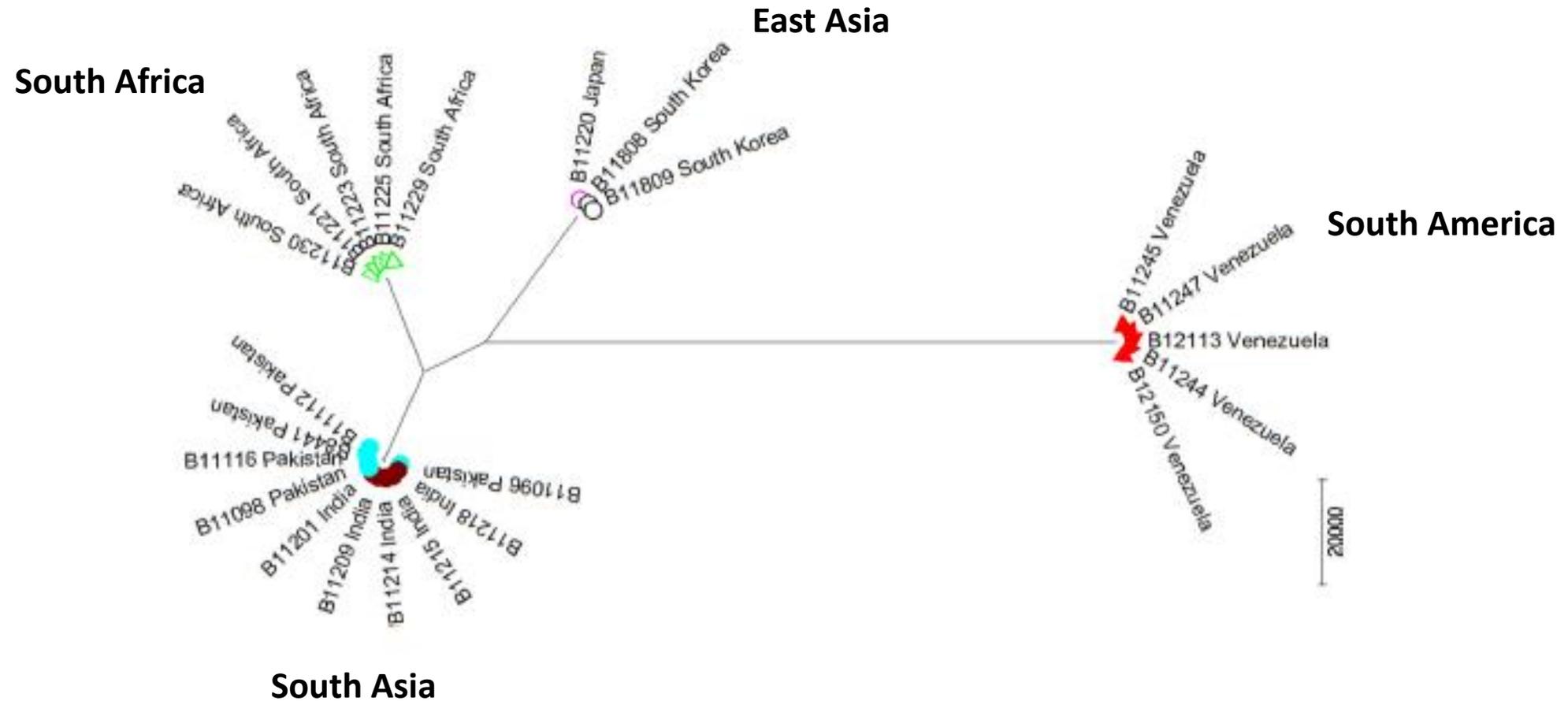
- **Work with laboratories, healthcare facilities and providers in your jurisdiction**
 - Prompt notification and reporting of **confirmed or suspect *C. auris*** cases
- **Facilitate submission of confirmed or suspect isolates to the MDHHS Bureau of Labs**
- **Enter case into MDSS and collect needed laboratory and clinical information**
- **Support implementation of Novel MDRO Containment Strategies**
 - Reduce transmission of *C. auris* in healthcare facilities
- **SHARP Unit can guide you through this process...**

Thank You

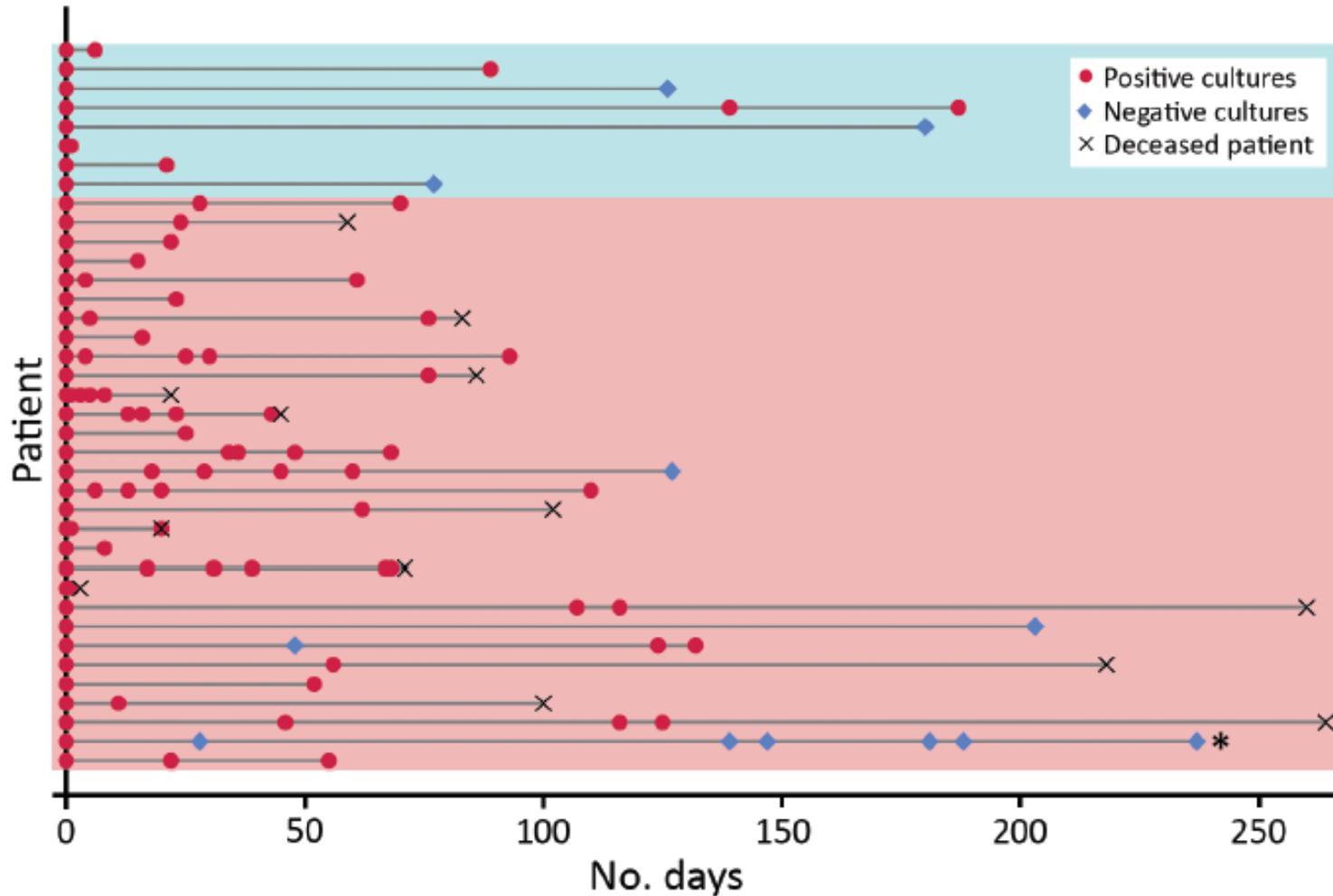
**Surveillance for Healthcare Associated and Resistant Pathogens (SHARP) Unit
Michigan Department of Health and Human Services (MDHHS)
(517) 335-8165**



Four Distinct Clades of *Candida auris*



Duration of Colonization Can Be Prolonged



Higher Colonization Rates in High-Acuity LTCF

- In NY, 572 patients screened in 19 facilities for *Candida auris* colonization
 - 61 (11%) were positive for *C. auris*
 - 19 (31%) admitted to acute care hospital
 - 42 (67%) resided at a long-term care facility
 - 40 (66%) were at a vSNF
- In Chicago, 1,364 patients screened in 20 facilities
 - 92 (6.7%) were positive for *C. auris*
 - Prevalence median (range):
 - vSNF - 7.7% (0-43.3%)
 - LTAC - 0% (0-14.3%)
 - Acute care - 0% (0-6.3%)
 - SNF - 0% (0-1.5%)

